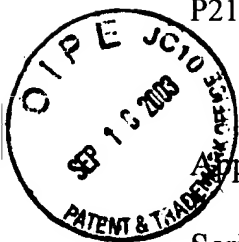


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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant : Arne HOLMGREN et al.

Group Art Unit : 1621

Serial No : 09/926,218
(National Stage of PCT/JP00/02076)

Examiner : Kumar

Filed : September 25, 2001
(International Filing Date March 31, 2000)

For : SUBSTRATE FOR THIOREDOXIN REDUCTASE

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

RECEIVED

SEP 12 2003

TECH CENTER 1600/2900

Sir:

In accordance with the duty of disclosure under 37 C.F.R. 1.56, 1.97, and 1.98, Applicants hereby brings the following additional information to the attention of the Examiner in charge of the above-identified application:

Engman et al., "Diaryl chalcogenides as selective inhibitors of thioredoxin reductase and potential antitumor agents", Anticancer Res., 1997 Nov-Dec;17(6D):4599-605 (Abstract)

Klotz et al., "Defenses against peroxynitrite: selenocompounds and flavonoids", Toxicol Lett., 2003 Apr 11;140-141:125-32 (Abstract).

Zhao et al., "A novel antioxidant mechanism of ebselen involving ebselen diselenide, a substrate of mammalian thioredoxin and thioredoxin reductase", J Biol Chem. 2002 Oct

18;277(42):39456-62.

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Zhao et al., "Ebselen: A substrate for human thioredoxin reductase strongly stimulating its hydroperoxide reductase activity and a superfast thioredoxin oxidant", Proc Natl Acad Sci U S A. 2002 Jun 25;99(13):8579-84.

Holmgren et al., "Antioxidant function of thioredoxin and glutaredoxin systems", Antioxid Redox Signal. 2000 Winter;2(4):811-20 (Abstract).

Sies et al., "Interaction of peroxynitrite with selenoproteins and glutathione peroxidase mimics", Free Radic Biol Med. 2000 May 15;28(10):1451-5 (Abstract).

Arteel et al., "Function of thioredoxin reductase as a peroxynitrite reductase using selenocystine or ebselen", Chem Res Toxicol. 1999 Mar;12(3):264-9 (Abstract).

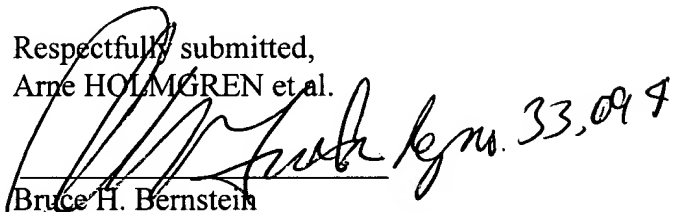
Copies of the above-noted documents are enclosed together with a duly completed Form PTO-1449. The Examiner is accordingly requested to consider each of these documents, and to make them of record in this application by initialing in the appropriate spaces on the Form PTO-1449. Applicants respectfully request that the Examiner include a copy of the initialed Form PTO-1449 with the next communication from the U.S. Patent and Trademark Office.

Applicants enclose a check including the amount of \$180.00 for consideration of this disclosure statement which is being submitted after a first Office Action on the merits, but prior to mailing of an action closing prosecution. If any fee is necessary for consideration of this disclosure statement, including any fee under 37 C.F.R. 1.17(p), this is authorization to charge Deposit Account No. 19-0089 any required fee.

P21480.A07

Should there be any questions, the Examiner is invited to contact the undersigned at the below listed telephone number.

Respectfully submitted,
Arne HOLMGREN et al.


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September 10, 2003
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